Appl. No. 09/492,028 Amdt. dated March 18, 2004 Reply to Office Action of October 1, 2003, and the Advisory Action mailed November 14, 2003

Amendments to the Specification:

Please replace paragraph on page 4 under the title SUMMARY OF THE INVENTION with the following amended paragraph:

-- The present invention demonstrates, for the first time, taste receptor cell specific expression of nucleic acids encoding G-protein alpha subunit. Specifically, the present invention identifies that $G\alpha 14$, a G-protein alpha subunit, is specifically and selectively expressed in taste receptor cells. This gene was found to be co-expressed with G-protein coupled taste receptors, GPCR-B3 and GPCR-B4 (see USSN 09/361,652, filed July 27, 1999 and USSN. 09/361,631, filed July 27, 1999, now U.S. Patent No. 6,383,778). These taste receptors have been previously shown to be expressed in topographically distinct subpopulations of taste receptor cells and taste buds. These receptors are specifically localized to the taste pore, and are distantly related to putative mammalian pheromone receptors. The present invention thus demonstrates that $G\alpha 14$ is specifically expressed in taste cells and further that it is co-expressed with GPCR-B3 and GPCR-B4 receptors in the different taste papillae. The G-protein alpha subunits that are specifically expressed in taste cells can thus be used, e.g., to screen for modulators of taste. The compounds identified by these assays would then be used by the food and pharmaceutical industries to customize taste, e.g., as additives to food or medicine so that the food or medicine tastes different to the subject who ingests it. For example, bitter medicines can be made to taste less bitter, and sweet substance can be enhanced.--

Please replace the paragraph starting on page 11, line 24, with the following amended paragraph:

--"TC-GPCR" refers to a G-protein coupled receptor that is specifically expressed in taste receptor cells such as foliate, fungiform, and circumvallate cells. Such taste cells can be identified because they express molecules such as Gustducin, a taste cell specific G-protein (McLaughin et al., Nature 357:563-569 (1992)). Taste receptor cells can also be identified on the basis of morphology (see, e.g., Roper, supra). Examples of TC-GPCR include GPCR-B3 and GPCR-B4 (see, e.g., Hoon et al., Cell 96:541-551 (1999); see also USSN 09/361,652, filed

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July 27, 1999 and USSN. 09/361,631, filed July 27, 1999, now U.S. Patent No. 6,383,778), herein incorporated by reference in their entirety). TC-GPCRs encode G-protein coupled receptors with seven transmembrane regions that have "G-protein coupled receptor activity," as described below, e.g., they bind to G-proteins in response to extracellular stimuli and promote production of second messengers such as IP₃, cAMP, and Ca²⁺ via stimulation of enzymes such as phospholipase C and adenylate cyclase (for a description of the structure and function of G-protein coupled receptors, *see*, *e.g.*, Fong, *supra*, and Baldwin, *supra*).--

Please replace the paragraph starting on page 32, line 25, with the following amended paragraph:

-- In a preferred embodiment, TC-Gα14 activity is measured by expressing TC-Gα14 in a heterologous cell with a TC-GPCR (see USSN 09/361,652, filed July 27, 1999 and USSN 09/361,631, filed July 27, 1999, now U.S. Patent No. 6,383,778). As shown in Example I below, TC-Gα14 is specifically expressed in taste receptor cells, and also co-expressed with GPCR-B3 and GPCR-B4, in different taste papillae. As described above, HEK-293 cells may be used as a heterologous host cell, and modulation of taste transduction is assayed by measuring changes in intracellular Ca²⁺ levels.--

Please replace the paragraph starting on page 59, line 24, with the following amended paragraph:

-- These experiments demonstrate that $G\alpha_{14}$ is specifically and selectively expressed in circumvallate, foliate and fungiform taste receptor cells of the tongue, as shown by *in situ* hybridization. Therefore, $G\alpha_{14}$ is a G alpha subunit that is specifically expressed in taste receptor cells. Furthermore, this gene is co-expressed with both GPCR-B3 and GPCR-B4 receptors in the different taste papillae (*see* USSN 09/361,652, filed July 27, 1999 and USSN. 09/361,631, filed July 27, 1999, now U.S. Patent No. 6,383,778).--